Attorney Docket No.: 960296.95360 Applicant: Ronald T. Raines

Application No.: 09/234,028 Filed: 01/20/1999

Date of Office Action: 12 DEC 2008 Group Art Unit: 1652

Examiner: Richard G. Hutson Date of Response: February 12, 2009

AMENDMENT TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

- (Currently amended) An oxidation-resistant engineered ribonuclease inhibitor (RI)
 variant selected from the group consisting of
- (i) [[an]] a RI variant differing from comprising the reference sequence of SEQ ID NO: 3, wherein the difference consists of except for an alanine substitution of at least one cysteine residue at position positions 95, 96, 329 or and 330; and
- (ii) [[an]] a RI variant differing from comprising the reference sequence of SEQ ID
 NO: 2, wherein the difference consists of except for an alanine substitution of at least one cysteine residue at position positions 324 or and 325,

the variant RI having a greater resistance to oxidation relative to the reference sequence for the variant RI and retaining its specificity and binding affinity to ribonuclease.

- (Cancelled)
- (Cancelled)
- 4. (Currently amended) The ribonuclease inhibitor variant <u>RI</u> of <u>elaimClaim</u> 1, wherein the <u>difference alanine substitution</u> inhibits the formation of a disulfide bond with an adjacent cysteine residue.

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- 5. (Currently amended) The ribonuclease inhibitor variant RI of elaim 18 Claim 1, wherein the variant RI is 10 to 15 fold more resistant to oxidative damage than a-native ribonuclease inhibitor an RI comprising the relevant reference sequence.
 - 6. (Cancelled)
- 7. (Currently amended) The ribonuclease inhibitor variant <u>RI</u> of elaim <u>Claim</u> 1, wherein the variant RI exhibits an *in vitro* inhibition of ribonucleolytic activity.
 - 8. (Cancelled)
- (Currently amended) The ribonuclease inhibitor variant RI of elaim Claim 18, having wherein the RI retains specificity and affinity of SEQ ID NO:3 for binding to angiogenin of the reference SEQ ID NO:3.
 - 10. 17. (Cancelled)
- 18. (Currently amended) An oxidation-resistant ribonuclease inhibitor (RI) variant differing from comprising the reference sequence of SEQ ID NO: 3, wherein the difference consists of except for an alanine residue at position one or more of positions 95, 96, 329 or and 330.
- (Currently amended) [[An]]The RI variant of Claim 18, wherein the residues at positions 329 and 330-beine are alanine residues.

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20. (Currently amended) An oxidation-resistant ribonuclease inhibitor (RI) variant differing from comprising the reference sequence of SEQ ID NO: 2, wherein the difference sensists of except for an alanine residue at position one or both of positions 324 or and 325.

- (Currently amended) [[An]]The RI variant of Claim 20, wherein the residues at positions 324 and 325-being are alanine residues.
- 22. (Currently amended) [[An]]The RI variant of Claim 18, wherein the residues at positions 95 and 96-being are alanine residues.
- 23. (Currently amended) [[An]]The RI variant of Claim 18, wherein the residues at positions 95, 96, 329 and 330-being are alanine residues.